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NOTICE OF ALLOWANCE AND FEE(S) DUE

58249 7590 06/25/2008 COOLEY GODWARD KRONISH LLP ATTN: Patent Group Suite 1100

777 - 6th Street, NW WASHINGTON, DC 20001 EXAMINER

MARVICH, MARIA

ART UNIT PAPER NUMBER

1633

DATE MAILED: 06/25/2008

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR ATTORNEY DOCK		CONFIRMATION NO.
10/629,453	07/29/2003	Jack D. Keene	RIBO-001/04US	5725
TITLE OF INVENTION: M	ETHODS FOR ISOLATING	G AND CHARACTERIZING ENDOGENOUS MRNA-PROTI	EIN (MRN77°C-2011)LEXES	

 APPLN. TYPE
 SMALL ENTITY
 ISSUE FEE DUE
 PUBLICATION FEE DUE
 PREV. PAID ISSUE FEE
 TOTAL FEES) DUE
 DATE DUE

 nonprovisional
 YES
 \$720
 \$300
 \$0
 \$1020
 09/25/2008

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

HOW TO REPLY TO THIS NOTICE:

I. Review the SMALL ENTITY status shown above.

If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:

A. If the status is the same, pay the TOTAL FEE(S) DUE shown above.

B. If the status above is to be removed, check box 5b on Part B Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

If the SMALL ENTITY is shown as NO:

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and I/2 the ISSUE FFE shown above.

II. PART B - FEE(S) TRANSMITTAL, or its equivalent, must be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be completed and an extra copy of the form should be submitted. If an equivalent of Part B is filed, a request to reapply a previously paid issue fee must be clearly made, and delays in processing may occur due to the difficulty in recognizing the paper as an equivalent of Part B.

III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

PART B - FEE(S) TRANSMITTAL

Complete and send this form, together with applicable fee(s), to: Mail Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450

or Fax (571)-273-2885

INSTRUCTIONS: This form should be used for transmitting the ISSUE DEE and DURI ICATION DEE (if required). Blocks 1 through 5 should be completed where

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777 - 6th Street, WASHINGTON								(Depositor's name)
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								(Date)
APPLICATION NO.	APPLICATION NO. FILING DATE		FIRST NAMED INVENTOR		ATTORNEY DOCKET NO. CONFIRMA		FIRMATION NO.	
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APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSU	E FEE	TOTAL FEE(S) DUE		DATE DUE
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MARVICE		I633	435-006000					
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PLEASE NOTE: Uni recordation as set forti (A) NAME OF ASSIG	less an assignee is ident h in 37 CFR 3.11. Comp GNEE	ified below, no assignee pletion of this form is NO	THE PATENT (print or ty data will appear on the p IT a substitute for filing an (B) RESIDENCE: (CITY	natent. If an assign assignment. A and STATE OR	COUNT	TRY)		
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NOTE: The Issue Fee and interest as shown by the i	d Publication Fee (if req records of the United Sta	uired) will not be accepte tes Patent and Trademark	d from anyone other than to Office.					
Authorized Signature				Date				
Typed or printed name	e			Registration !	No			
This collection of inform an application. Confident submitting the complete this form and/or suggesti Box 1450, Alexandria, V Alexandria, Virginia 223	ation is required by 37 C tiality is governed by 35 d application form to the ons for reducing this but firginia 22313-1450. DC 13-1450.	FR 1.311. The informatis U.S.C. 122 and 37 CFR USPTO. Time will vary rden, should be sent to th O NOT SEND FEES OR	on is required to obtain or 1.14. This collection is es depending upon the indi- e Chief Information Offic COMPLETED FORMS T	retain a benefit by timated to take 12 vidual case. Any co er, U.S. Patent and O THIS ADDRES	the pub minuter omment Trader S. SEN	lic which is to file (and s to complete, including s on the amount of tit mark Office, U.S. Dep. D TO: Commissioner	i by the ig gathe ne you artment for Pate	USPTO to process) ring, preparing, and require to complete of Commerce, P.O. rnts, P.O. Box 1450,

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10/629,453	07/29/2003	Jack D. Keene	RIBO-001/04US 308729-2034	5725	
58249 75	590 06/25/2008		EXAM	IINER	
COOLEY GODWARD KRONISH LLP MARVICH, MARIA			I, MARIA		
ATTN: Patent Group			ART UNIT	PAPER NUMBER	
Suite 1100 1633					
777 - 6th Street, NW		DATE MAILED: 06/25/2008			

Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 0 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 0 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 or (571)-272-4200.

Notice of Allowability

Application No.	Applicant(s)		
10/629,453	KEENE ET AL.		
Examiner	Art Unit		
MADIA R. MADVICH	1622		

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address--All claims being allowable, PROSECUTION ON THE MERITS IS (OR REMAINS) CLOSED in this application. If not included herewith (or previously mailed), a Notice of Allowance (PTOL-85) or other appropriate communication will be mailed in due course. THIS NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIGHTS. This application is subject to withdrawal from issue at the initiative

- of the Office or upon petition by the applicant. See 37 CFR 1.313 and MPEP 1308.
- This communication is responsive to an after final filed 5/19/08.
- The allowed claim(s) is/are 30-38 and 40-62.
- 3. Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
 - b) ☐ Some* c) ☐ None of the:
 - 1. T Certified copies of the priority documents have been received.
 - 2. Certified copies of the priority documents have been received in Application No.
 - 3. Copies of the certified copies of the priority documents have been received in this national stage application from the International Bureau (PCT Rule 17.2(a)).
 - * Certified copies not received: _____.

Applicant has THREE MONTHS FROM THE "MAILING DATE" of this communication to file a reply complying with the requirements noted below. Failure to timely comply will result in ABANDONMENT of this application. THIS THREE-MONTH PERIOD IS NOT EXTENDABLE.

- 4. A SUBSTITUTE OATH OR DECLARATION must be submitted. Note the attached EXAMINER'S AMENDMENT or NOTICE OF INFORMAL PATENT APPLICATION (PTO-152) which gives reason(s) why the oath or declaration is deficient.
- CORRECTED DRAWINGS (as "replacement sheets") must be submitted.
 - (a) Including changes required by the Notice of Draftsperson's Patent Drawing Review (PTO-948) attached
 - 1) hereto or 2) to Paper No./Mail Date
 - (b) including changes required by the attached Examiner's Amendment / Comment or in the Office action of

Identifying indicia such as the application number (see 37 CFR 1.84(c)) should be written on the drawings in the front (not the back) of each sheet. Replacement sheet(s) should be labeled as such in the header according to 37 CFR 1.121(d).

6.

DEPOSIT OF and/or INFORMATION about the deposit of BIOLOGICAL MATERIAL must be submitted. Note the attached Examiner's comment regarding REQUIREMENT FOR THE DEPOSIT OF BIOLOGICAL MATERIAL.

Attachment(s)

- 1. | Notice of References Cited (PTO-892)
- 2. Notice of Draftperson's Patent Drawing Review (PTO-948)
- Information Disclosure Statements (PTO/SB/08).
- Paper No./Mail Date 4. ☐ Examiner's Comment Regarding Requirement for Deposit of Biological Material
- 5. Notice of Informal Patent Application Interview Summary (PTO-413), Paper No./Mail Date 7. X Examiner's Amendment/Comment
- 8. X Examiner's Statement of Reasons for Allowance
- Other .

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DETAILED ACTION

This office action is in response to an amendment filed 10/1/04.

EXAMINER'S AMENDMENT

An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with Konstatin Linnik on 6/19/08.

The application has been amended as follows:

IN THE SPECIFICATION:

The priority data has been amended to read: In the specification, page 1, line 1,

-- RELATED APPLICATIONS

This application is a divisional of copending U.S. Patent Application Serial No. 09/750,401 filed December 12, 2001, now Patent No. 6,635,422, which claims priority to U.S. Provisional Patent Application Serial No. 60/173,338 filed December 28, 1999. --

IN THE CLAIMS:

30. (Currently amended) A method of identifying an endogenous mRNA subset[[s]] in a cell comprising an mRNA-protein (mRNP) complex, comprising the steps of:

- (a) lysing a the cell comprising an mRNA-protein (mRNP) complex to produce a lysate;
- (b) contacting the lysate with an antibody that specifically binds at least one component of the mRNP complex;
- (c) partitioning the mRNP complex by eapturing $\underline{binding}$ the antibody on \underline{to} a solid support; (d) removing the eaptured \underline{bound} mRNP complex from the lysate; and
- (e) identifying a plurality of mRNAs from the mRNP complex without amplifying the mRNAs by PCR, wherein the identified mRNAs are encoded by a plurality of <u>distinct genes and</u> wherein the identified mRNAs form the endogenous mRNA subset in the cell.

In claim 37 insert the phrase --at least one-- prior to "the component of the" and delete the word "captured" following this phase and insert the word --bound--.

In claim 43 insert the article --a-- prior to the term "plant cell".

In claim 45 delete the number "30" following the phrase "method of claim" and insert the number --37--.

48. (Currently amended): A method of identifying an endogenous mRNA subset[[8]] in a cell, comprising the steps of:

(a) expressing an epitope-tagged RNA-binding protein or an epitope-tagged RNAassociated protein (RAP) in a the cell, thereby forming an mRNP complex;

- (b) lysing the cell to produce a lysate;
- (c) partitioning the mRNP complex by eapturing <u>binding</u> the RNA binding protein or the RAP on to a solid support;
 - (d) removing the eaptured bound mRNP complex from the lysate; and
- (e) identifying a plurality of mRNAs from the mRNP complex without amplifying the mRNAs by PCR, wherein the identified mRNAs are encoded by a plurality of <u>distinct genes</u> and wherein the identified mRNAs form the endogenous mRNA subset in the cell.
- 58. (Currently amended) The method of claim 48, wherein the mRNP complex is contacted with the epitope-tagged RNA-binding protein is expressed in the cell.

In claim 62 insert the article --a-- prior to the term "plant cell".

EXAMINER'S AMENDMENT

The following is an examiner's statement of reasons for allowance. The instant application teaches methods of *in vivo* selection of physically clustered mRNA subsets associated with RNA binding or RNA associated proteins. The specification teaches that many of these mRNA subsets are also functionally associated with particular cellular conditions and hence monitoring alterations in the subsets following compound treatment, aging, differentiation, development and disease state may be characteristic of particular diseases. The closest prior art

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is Giordano et al (US 5,859,227 and WO 98/37422), which teaches method of identifying compounds that modulate interactions between RNA binding proteins and RNA molecules in a sample such as a cell or tissue extract. The methods of Giordano et al measure the interaction by use of filter binding as well as gel shift mobility. In contrast, the instant method involves isolating cellular mRNP complexes by immunoprecipitation such that associated subsets of RNA can be identified without further steps of amplification or iterative selection. While methods of immunoprecipitation of RNA binding proteins are known in the art, the instant invention recognized that immuno- precipitated RNA binding proteins comprises subsets of RNA that could be removed and directly identified and quantitated. Hence, the instant method advances the prior art by providing a method of measuring changes in functionally critical subsets of mRNA in vivo by isolation of associated RNA binding or associated proteins and direct analysis of the associated mRNAs.

Conclusion

Any inquiry concerning this communication or earlier communications from the examiner should be directed to MARIA B. MARVICH whose telephone number is (571)272-0774. The examiner can normally be reached on M-F (7:00-4:00).

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Joseph Woitach, PhD can be reached on (571)-272-0739. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

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Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated

information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

Maria B Marvich, PhD Primary Examiner Art Unit 1633

/Maria B Marvich, PhD/ Primary Examiner, Art Unit 1633